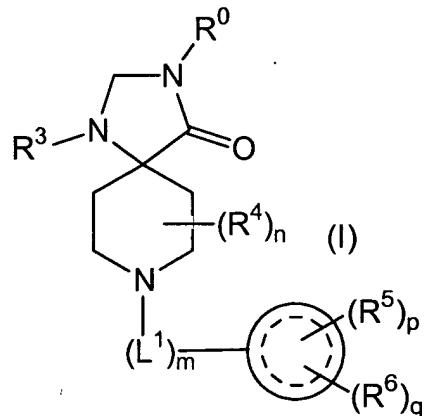


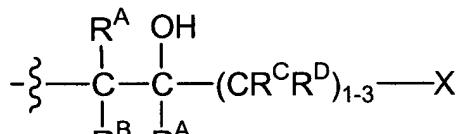
In the Claims:

This listing of Claims will replace all prior versions, and listings, of Claims in the application.

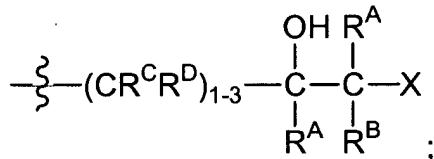
1. (Currently Amended) A compound of the formula (I)



wherein



R^0 is selected from the group consisting of and



each R^A and R^B is independently selected from the group consisting of hydrogen and C₁₋₄alkyl;

each R^C and R^D is independently selected from the group consisting of hydrogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, nitro, cyano, $N(R^E)_2$, aryl, arC₁₋₄alkyl, heteroaryl or heterocycloalkyl; wherein the aryl, arC₁₋₄alkyl, heteroaryl or heterocycloalkyl substituent is optionally substituted with one or more substituents independently selected from hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, nitro, cyano or $N(R^E)_2$;

each R^E is independently selected from the group consisting of hydrogen and C₁₋₄alkyl;

X is selected from the group consisting of $\text{-NR}^1\text{R}^2$, $\text{-C(O)-NR}^1\text{R}^2$, $\text{-NR}^1\text{-C(O)-R}^2$, -OR^1 , -SR^1 , -SOR^1 , $\text{-SO}_2\text{R}^1$, $\text{-S-(C}_{2-4}\text{alkyl)-NR}^1\text{R}^2$, $\text{-S-(C}_{2-4}\text{alkyl)-C(O)O-C(CH}_3)_3$, $\text{-SO-(C}_{1-4}\text{alkyl)-NR}^1\text{R}^2$ and $\text{-SO}_2\text{(C}_{1-4}\text{alkyl)-NR}^1\text{R}^2$; wherein the alkyl portion of the $\text{-S-(C}_{2-4}\text{alkyl)-NR}^1\text{R}^2$, $\text{-SO-(C}_{1-4}\text{alkyl)-NR}^1\text{R}^2$ or $\text{-SO}_2\text{(C}_{1-4}\text{alkyl)-NR}^1\text{R}^2$ group is optionally substituted with one or more substituents independently selected from carboxy, hydroxy, hydroxyC₁₋₄alkyl, C₁₋₄alkyl, C₁₋₄alkoxycarbonyl or CONR¹R²;

each R¹ and R² is independently selected from the group consisting of hydrogen, C₁₋₈alkyl, C₁₋₈alkoxy, C₁₋₈alkoxycarbonyl, cycloalkyl, cycloalkyl-C₁₋₄alkyl, partially unsaturated carbocyl, partially unsaturated carbocycl-C₁₋₄alkyl, aryl, arC₁₋₄alkyl, arC₁₋₄alkoxy, heteroaryl, heteroaryl-C₁₋₄alkyl, heterocycloalkyl, heterocycloalkyl-C₁₋₄alkyl, -C(O)-C₁₋₆alkyl, -C(O)-aryl, -C(O)-arC₁₋₄alkyl, -C(O)-heteroaryl, -C(O)-heterocycloalkyl, -C(O)O-cycloalkyl and -C(O)O-aryl, -C(O)O-arC₁₋₄alkyl, and -C(O)O-(partially unsaturated carbocycl), -C(O)O-heteroaryl, -C(O)O-heterocycloalkyl; wherein the C₁₋₈alkyl, cycloalkyl, partially unsaturated carbocycl, aryl, or arC₁₋₈alkyl, heteroaryl or heterocycloalkyl group, whether alone or part of a substituent group, is optionally substituted with one or more substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, trifluoromethyl, trifluoromethoxy, nitro, cyano, -C(O)-C₁₋₄alkyl, C₁₋₄alkoxycarbonyl, N(R^E)₂, N(R^E)₂-C₁₋₄alkyl, N(R^E)-C(O)C(CH₃)₃, -C₁₋₄alkyl-N(R^E)-C(O)O-C₁₋₄alkyl and -N(R^E)-C(O)O-C₁₋₄alkyl, aryl, aryloxy, cycloalkyl, heteroaryl, aryl substituted heteroarylaminosulfonyl or C₁₋₆alkylthio;

alternatively when R¹ and R² are both bound to the same nitrogen atom, R¹ and R² are taken together with the nitrogen atom to which they are bound to form a heteroaryl or heterocycloalkyl group; wherein the heteroaryl or heterocycloalkyl group is optionally substituted with one or more substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, hydroxy substituted C₁₋₄alkyl, C₁₋₄alkoxy, C₁₋₄alkoxycarbonyl, trifluoromethyl, trifluoromethoxy, nitro, cyano, N(R^E)₂, aryl, arC₁₋₄alkyl, heteroaryl, heterocycloalkyl, di(C₁₋₆)alkylamine-carbonyl, C₁₋₄alkoxycarbonyl-N(R^E) or arylamine-C₁₋₄alkyl; wherein the aryl, arC₁₋₄alkyl, heteroaryl or heterocycloalkyl substituent is optionally further substituted with one or more substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, trifluoromethyl, trifluoromethoxy, nitro, cyano,

~~N(R^E)₂, phenyl or substituted phenyl; wherein the substituents on the phenyl are one or more independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, trifluoromethyl, trifluoromethoxy, nitro, cyano or N(R^E)₂;~~

R³ is selected from the group consisting of aryl, arC₁₋₆alkyl and heteroaryl; wherein the aryl, arC₁₋₆alkyl or heteroaryl group is optionally substituted with one or more substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, trifluoromethyl, trifluoromethoxy, nitro, cyano or N(R^E)₂;

n is an integer from 0 to 2;

R⁴ is selected from the group consisting of hydroxy, C₁₋₄alkyl and hydroxy substituted C₁₋₄alkyl;

m is an integer from 0 to 1;

L¹ is selected from the group consisting of C₁₋₆alkyl and C₃₋₆alkenyl; wherein the double bond of the C₃₋₆alkenyl group is at least one carbon atom removed from the attachment point to the N atom; and wherein the C₁₋₆alkyl or C₃₋₆alkenyl group is optionally substituted with one to two substituents independently selected from hydroxy, fluoro, C₁₋₆alkyl, fluorinated C₁₋₆alkyl or C₁₋₆alkoxy;



is selected from the group consisting of cycloalkyl, partially unsaturated carbocyclyl, aryl, heteroaryl and heterocycloalkyl phenyl, naphthyl and acenaphthyl;

p is an integer from 0 to 5;

R⁵ is selected from the group consisting of hydroxy, carboxy, halogen, C₁₋₆alkyl, hydroxy substituted C₁₋₆alkyl, C₁₋₆alkoxy, nitro, cyano, NR¹R², trifluoromethyl, trifluoromethoxy, C₁₋₄alkoxycarbonyl, -SO-NR¹R², -SO₂-NR¹R² and -C(O)-NR¹R²;

q is an integer from 0 to 1;

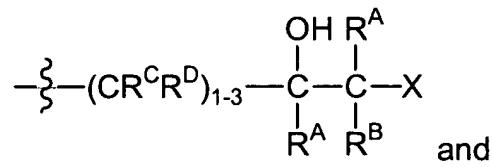
R⁶ is selected from the group consisting of -(L²)₀₋₁-R⁷;

L² is selected from the group consisting of -C₁₋₆alkyl-, -C₂₋₄alkenyl-, -C₂₋₆alkynyl-, -O-, -S-, -NH-, -N(C₁₋₄alkyl)-, -C₁₋₆alkyl-O-, -C₁₋₆alkyl-S-, -O-C₁₋₆alkyl-, -S-C₁₋₆alkyl-, -O-C₂₋₆alkyl-O-, -S-C₂₋₆alkyl-S-, -SO₂-, -SO₂NH-, -SO₂N(C₁₋₄alkyl)-, -NH-SO₂-, -N(C₁₋₄alkyl)-SO₂-, -C(O)-O- and -O-C(O)-;

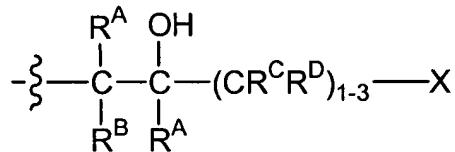
R^7 is selected from the group consisting of aryl, partially unsaturated carbocyclyl, cycloalkyl, heteroaryl and heterocycloalkyl; wherein the aryl, partially unsaturated carbocyclyl, cycloalkyl, heteroaryl or heterocycloalkyl group is optionally substituted with one or more substituents independently selected from hydroxy, carboxy, halogen, C_{1-6} alkyl, C_{1-6} alkoxy, nitro, cyano, $N(R^E)_2$, trifluoromethyl, trifluoromethoxy, C_{1-4} alkoxycarbonyl, $-SO_2-N(R^E)_2$ and $-C(O)-N(R^E)_2$;

or a pharmaceutically acceptable salt thereof.

2. (Currently Amended) A compound as in Claim 1 wherein



R^0 is selected from the group consisting of



each R^C and R^D is independently selected from hydrogen, C_{1-4} alkyl, C_{1-4} alkoxy, hydroxy, carboxy or aryl; wherein the aryl is optionally substituted with one to two substituents independently selected from hydroxy, carboxy, C_{1-4} alkyl, C_{1-4} alkoxy, nitro, cyano or $N(R^E)_2$;

~~X is selected from the group consisting of $-NR^1R^2$, $-C(O)-NR^1R^2$, $-NR^1-C(O)-R^2$, OR^1 , SR^1 , $SO-R^1$, SO_2-R^1 , $S-(C_{2-4}\text{alkyl})NR^1R^2$, $S-(C_{2-4}\text{alkyl})NR^1-C(O)OC(CH_3)_3$, $SO-(C_{1-4}\text{alkyl})NR^1R^2$ and $SO_2-(C_{1-4}\text{alkyl})NR^1R^2$; wherein the alkyl portion of the $S-(C_{2-4}\text{alkyl})NR^1R^2$, $SO-(C_{1-4}\text{alkyl})NR^1R^2$ or $SO_2-(C_{1-4}\text{alkyl})NR^1R^2$ group is optionally substituted with one to two substituents independently selected from C_{1-4} alkyl, hydroxy C_{1-4} alkyl, C_{1-4} alkoxycarbonyl or carboxy;~~

R^1 is selected from the group consisting of hydrogen, C_{1-4} alkyl, C_{1-4} alkoxy, C_{1-4} alkoxycarbonyl, aryl, ar C_{1-4} alkyl, ar C_{1-4} alkyloxy, heteroaryl, heteroaryl-alkyl, heterocycloalkyl, heterocycloalkyl-alkyl, cycloalkyl-alkyl, and $C(O)-C_{1-4}$ alkyl and $C(O)$ -heteroaryl;

wherein the C₁₋₄alkyl, aryl, arC₁₋₄alkyl, heteroaryl, heterocycloalkyl or cycloalkyl group, whether alone or part of a substituent group, is optionally substituted with one to three substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, C₁₋₄alkoxycarbonyl, N(R^E)₂, N(R^E)₂-C₁₋₄alkyl, N(R^E)-C(O)OC(CH₃)₃, nitro, trifluoromethyl, trifluoromethoxy, phenyl, phenoxy, heteroaryl, cycloalkyl, 1-phenyl-pyrazol-2-yl-aminosulfonyl or C₁₋₄alkylthio;

R² is selected from the group consisting of hydrogen, C₁₋₄alkyl, C₁₋₄alkoxy, cycloalkyl, cycloalkyl-C₁₋₄alkyl, aryl, arC₁₋₄alkyl, arC₁₋₄alkyloxy, partially unsaturated carbocyclyl, partially unsaturated carbocyclyl-C₁₋₄alkyl, heteroaryl, heteroaryl-C₁₋₄alkyl, heterocycloalkyl, heterocycloalkyl-C₁₋₄alkyl, -C(O)-C₁₋₄alkyl, -C(O)-aryl, -C(O)-arC₁₋₄alkyl, -C(O)-heteroaryl, -C(O)-heterocycloalkyl, -C(O)O-cycloalkyl and -C(OO)-C₁₋₄alkyl;

wherein the C₁₋₄alkyl, aryl, arC₁₋₄alkyl, partially unsaturated carbocyclyl, heteroaryl, heterocycloalkyl or cycloalkyl group, whether alone or part of a substituent group, is optionally substituted with one to three substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, C₁₋₄alkoxycarbonyl, N(R^E)₂, N(R^E)₂-C₁₋₄alkyl, (CH₃)₃COC(O)-N(R^E)-C₁₋₄alkyl, nitro, cyano, trifluoromethyl, trifluoromethoxy, phenyl, phenoxy, heteroaryl, cycloalkyl, 1-phenyl substituted heteroaryl-aminosulfonyl, -C(O)-C₁₋₄alkyl or C₁₋₄alkylthio;

alternatively when R¹ and R² are both bound to the same nitrogen atom, R¹ and R² are taken together with the nitrogen atom to which they are bound to form a heteroaryl or heterocycloalkyl group;

wherein the heteroaryl or heterocycloalkyl is optionally substituted with one to three substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, hydroxy substituted C₁₋₄alkyl, C₁₋₄alkoxy, C₁₋₄alkoxycarbonyl, trifluoromethyl, trifluoromethoxy, nitro, cyano, N(R^E)₂, phenyl, arC₁₋₄alkyl, heterocycloalkyl, di(C₁₋₄alkyl)amino carbonyl, C₁₋₄alkoxycarbonylamino or phenylamino-C₁₋₄alkyl;

wherein the phenyl or arC₁₋₄alkyl substituent on the heteroaryl or heterocycloalkyl group is optionally substituted with one or two substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, trifluoromethyl, trifluoromethoxy, nitro,

~~cyano, $N(R^E)_2$ or substituted phenyl; wherein the substituents on the phenyl are one to three independently selected from halogen;~~

~~R³ is selected from the group consisting of aryl and arC₁₋₄alkyl; wherein the aryl or arC₁₋₄alkyl group is optionally substituted with one to three substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, trifluoromethyl, trifluoromethoxy, nitro, cyano or $N(R^E)_2$;~~

n is an integer from 0 to 1;

L¹ is C₁₋₄alkyl; wherein the C₁₋₄alkyl group is optionally substituted with one to two substituents independently selected from hydroxy, fluoro, C₁₋₄alkyl, fluorinated C₁₋₄alkyl or C₁₋₄alkoxy;

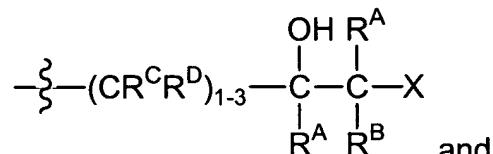
R⁵ is selected from the group consisting of hydroxy, carboxy, halogen, C₁₋₄alkyl, C₁₋₄alkoxy, nitro, cyano, $N(R^E)_2$, trifluoromethyl, trifluoromethoxy, C₁₋₄alkoxycarbonyl, -SO- $N(R^E)_2$, -SO₂- $N(R^E)_2$ and -C(O)- $N(R^E)_2$;

L² is selected from the group consisting of -C₁₋₄alkyl-, -O-, -S-, -N(R^E)-, -C(O)O- and -O-C(O)-;

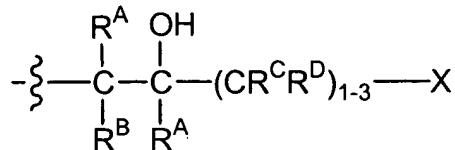
R⁷ is selected from the group consisting of cycloalkyl, aryl, heteroaryl and heterocycloalkyl; wherein the aryl, heteroaryl or heterocycloalkyl group is optionally substituted with one to two substituents independently selected from hydroxy, carboxy, halogen, C₁₋₄alkyl, C₁₋₄alkoxy, nitro, cyano, $N(R^E)_2$, trifluoromethyl, trifluoromethoxy or C₁₋₄alkoxycarbonyl;

or a pharmaceutically acceptable salt thereof.

3. (Currently Amended) A compound as in Claim 2 wherein



R⁰ is selected from the group consisting of



each R^A, R^B, R^C and R^D is hydrogen;

~~X is selected from the group consisting of -NR¹R², -OR¹, -SR¹, -S-(C₂-alkyl)-NR¹R² and -S-(C₂-alkyl)-NR¹-C(O)O-C(CH₃)₃; wherein the alkyl portion of the -S-(C₂-alkyl)-NR¹R² or -S-(C₁₋₄alkyl)-NR¹-C(O)O-C(CH₃)₃ group is optionally substituted with a carboxy or C₁₋₄alkoxycarbonyl group;~~

~~R¹ is selected from the group consisting of hydrogen, C₁₋₄alkyl, C₁₋₄alkoxycarbonyl, arC₁₋₄alkyl and C(O)-C₁₋₄alkyl;~~

~~wherein the C₁₋₄alkyl or aryl group, whether alone or part of a substituent group, is optionally substituted with one to two substituents independently selected from carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, C₁₋₄alkoxycarbonyl, N(R^E)₂ or N(R^E)-C(O)OC(CH₃)₃;~~

~~R² is selected from the group consisting of hydrogen, C₁₋₄alkyl, C₁₋₄alkoxy, cycloalkyl, aryl, arC₁₋₄alkyl, arC₁₋₄alkyloxy, partially unsaturated carbocyclyl, partially unsaturated carbocyclyl-C₁₋₄alkyl, heteroaryl, heteroaryl-C₁₋₄alkyl, heterocycloalkyl, heterocycloalkyl-C₁₋₄alkyl, cycloalkyl-C₁₋₄alkyl, -C(O)arC₁₋₄alkyl, -C(O)-heteroaryl, -C(OO)-cycloalkyl and -C(O)O-C₁₋₄alkyl;~~

~~wherein the C₁₋₄alkyl, aryl, arC₁₋₄alkyl, partially unsaturated carbocyclyl, heteroaryl, heterocycloalkyl or cycloalkyl group, whether alone or part of a substituent group, is optionally substituted with one to three substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, C₁₋₄alkoxycarbonyl, N(R^E)₂, N(R^E)₂-C₁₋₄alkyl, (CH₃)₃CO-C(O)-N(R^E)-C₁₋₄alkyl, nitro, trifluoromethyl, trifluoromethoxy, phenyl, phenoxy, heteroaryl, cycloalkyl, 1-phenyl-pyrazol-2-yl-aminosulfonyl or C₁₋₄alkylthio;~~

~~alternatively when R¹ and R² are both bound to the same nitrogen atom, R¹ and R² are taken together with the nitrogen atom to which they are bound to form a group selected from heterocycloalkyl and heteroaryl;~~

~~wherein the heteroaryl or heterocycloalkyl is optionally substituted with one or more substituents independently selected from hydroxy, carboxy, C₁₋₄alkyl, hydroxy substituted C₁₋₄alkyl, C₁₋₄alkoxy, phenyl, arC₁₋₄alkyl, heterocycloalkyl, C₁₋₄alkoxycarbonyl, amino, C₁₋₄alkylamino, di(C₁₋₄alkyl)amino, di(C₁₋₄alkyl)amino-carbonyl, t-butoxycarbonylamino or phenylamino-C₁₋₄alkyl;~~

~~wherein the phenyl or arC₁₋₄alkyl substituent is optionally substituted with one or two substituents independently selected from chloro, trifluoromethyl or chlorophenyl;~~

R³ is aryl; wherein the aryl group is optionally substituted with one or more substituents independently selected from halogen;

n is 0;

L¹ is C₁₋₄alkyl;

R⁵ is selected from the group consisting of halogen, C₁₋₄alkyl and trifluoromethyl;

R⁶ is -(L²)₀-R⁷;

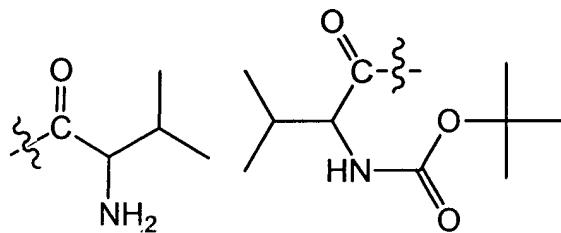
R⁷ is selected from the group consisting of aryl and heteroaryl;
or a pharmaceutically acceptable salt thereof.

4. (Currently Amended) A compound as in Claim 3 wherein

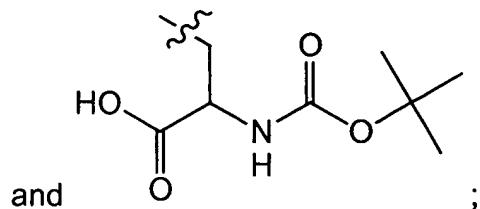
R⁰ is selected from the group consisting of -CH₂-CH(OH)-CH₂-X and -CH₂-CH₂-CH(OH)-CH₂-X;

X is selected from the group consisting of -NR¹R², -OR⁴, -SR⁴, -S-CH₂CH(CO₂H)-NH-C(O)-CH₃ and -S-CH₂CH(CO₂H)-NH-C(O)O-C(CH₃)₃;

R¹ is selected from the group consisting of hydrogen, methyl, ethyl, n-propyl, n-butyl, t-butyl, amino-n-propyl, dimethylaminoethyl, benzyl, phenylethyl, 4-methyl-benzyl,

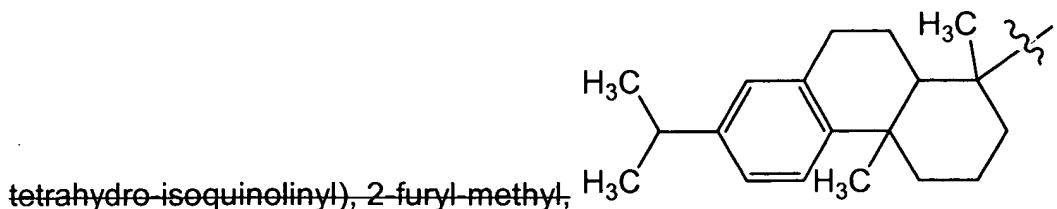


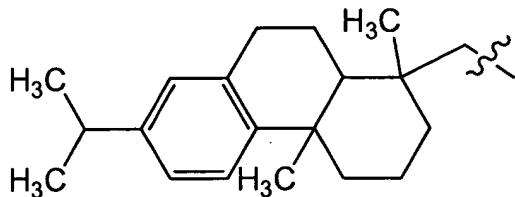
, 2-(3,4-dimethoxy-phenyl)ethyl, 3-methyl-phenyl, ethoxy-carbonyl-methyl, 2-amino-2-methoxycarbonyl-ethyl, t-butoxycarbonyl



R² is selected from the group consisting of hydrogen, methyl, methoxy, ethyl, carboxy-methyl, ethoxycarbonylmethyl, 2,2,2,-trifluoroethyl, ethoxy, dimethylaminoethyl, t-butoxycarbonylamino-ethyl, n-butyl, t-butyl, n-propyl, 3-hydroxy-n-propyl, 3-methoxy-n-

propyl, methylamino-n-propyl, dimethylamino-n-propyl, di(n-butyl)amino-n-propyl, t-butoxycarbonylamino-n-propyl, 3-phenyl-n-propyl, 3-(2-pyridyl)-n-propyl, t-butoxycarbonyl, cyclopropyl, phenyl, 4-fluorophenyl, 4-methylphenyl, 3,4-dimethoxyphenyl, 2-aminophenyl, 4-biphenyl, 2-ethoxyphenyl, 4-((1-phenyl-pyrazol-2-yl)-aminosulfonyl)-phenyl, 4-cyclohexylphenyl, 4-(aminoethyl)phenyl, 4-(t-butoxycarbonylamino-ethyl)-phenyl, -CH(CH₃)-phenyl, benzyl, benzyloxy, 2-methylbenzyl, 3-methylbenzyl, 4-methylbenzyl, 2-methoxybenzyl, 3-methoxybenzyl, 4-methoxybenzyl, 2-ethoxybenzyl, 3-ethoxybenzyl, 2-bromobenzyl, 3-bromobenzyl, 4-bromobenzyl, 3-chlorobenzyl, 4-chlorobenzyl), 3-iodobenzyl, 2-fluorobenzyl, 3-fluorobenzyl, 4-fluorobenzyl, 2-trifluoromethylbenzyl, 3-trifluoromethylbenzyl, 4-trifluoromethylbenzyl, 4-trifluoromethoxybenzyl, 4-methoxycarbonylbenzyl, 2,3-dimethoxybenzyl, 2,4-dichlorobenzyl, 3,4-dichlorobenzyl, 2,4-difluorobenzyl, 2,5-difluorobenzyl, 3,4-difluorobenzyl, 3,4,5-trimethoxybenzyl, 2,4,6-trimethoxybenzyl, 4-carboxybenzyl, 3-nitrobenzyl, 4-nitrobenzyl, 2,4-dimethoxybenzyl, 3,4-dimethoxybenzyl, 3,5-dimethoxybenzyl, 3,4-difluorobenzyl, 3,5-di(trifluoromethyl)benzyl, 4-(dimethylamino)benzyl, 2-phenylethyl, 2-(4-bromophenyl)ethyl, 2-(3-methoxyphenyl)ethyl, 2-(4-methoxyphenyl)ethyl, 2-(3,4-dimethoxyphenyl)ethyl, 2-(2-nitro-4,5-dimethoxy-phenyl)ethyl, 3-(4-morpholinyl)-n-propyl, 2-(4-morpholinyl)ethyl, 2-(4-imidazolyl)ethyl, 1-adamantanyl, 1-adamantanyl-methyl, (2,5-dimethoxy-2,5-dihydro-fur-2-yl)methyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, 2-pyridyl-methyl, 3-pyridyl-methyl, 4-pyridyl-methyl, 2-(3,4-dimethyl-pyridyl), 2-(5-bromopyridyl), 2-(4,6-dimethyl-pyridyl), 2-(5-methyl-pyridyl), 3-(6-methoxy-pyridyl), 6-methylthio-2-pyridyl-carbonyl, thienyl-methyl, 2-thienylethyl, 4-pyridinyl, 1-naphthyl, 1-naphthyl-methyl, 1-(3,4-methylenedioxyphe-nyl)methyl, 2-(3,4-methylenedioxyphe-nyl)ethyl, 1-phenyl-2-(t-butoxycarbonyl)ethyl, -C(O)-C(OCH₃)(CF₃)-phenyl, -C(O)O-(2-isopropyl-5-methyl-cyclohexyl), 1-(4-ethoxycarbonyl-piperidinyl), 2-(3H-imidazol-4-yl)ethyl, 2-(1,2,3,4-tetrahydro-isoquinolinyl), 2-furyl-methyl,



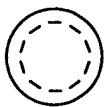


, 2S-hydroxy-S-cyclopentyl-methyl, 2S-hydroxy-S-cyclohexyl-methyl, 2S-hydroxy-S-cycloheptyl-methyl, 2-phenoxy-ethyl, 2-(2-pyridyl)-ethyl, 2-(6-fluoro-2-indolyl)ethyl and 2-phenyl-cyclopropyl;

alternatively when R¹ and R² are both bound to the same nitrogen atom, R¹ and R² are taken together with the nitrogen atom to which they are bound to form a group selected from 1-morpholinyl, 1-(4-(3-trifluoromethyl-phenyl)-piperazinyl), 1-(4-piperidinyl-piperidinyl), 1-(4-pyrrolidinyl-piperidinyl), 1-(4-phenyl-piperidinyl), 1-(3-hydroxy-piperidinyl), 1-(4-hydroxy-piperidinyl), 1-(3-hydroxymethyl-piperidinyl), 1-(3,5-dimethyl-piperidinyl), 1-(4-dimethylamino-piperidinyl), 1-(4-(3,4-methylenedioxyphenylmethyl)-piperazinyl), 1-(3-(diethylaminocarbonyl)-piperidinyl), 1-(4-t-butoxycarbonylamino-piperidinyl), 1-(2,3-dihydro-1H-pyrrolyl), 1-(4-[(4-chlorophenyl)-phenyl-methyl]-piperazinyl), 2-(1,2,3,4-tetrahydro-6,7-dimethoxy-isoquinolinyl), 1-(4-t-butoxycarbonyl-piperazinyl), 2-(1,2,3,4-tetrahydro-6,7-dimethoxy-isoquinolinyl), 4-(2,6-dimethyl-morpholinyl), 1-(4-benzyl-piperazinyl), 1-pyrrolidinyl, 1-(2,3-dihydro-pyrrolidinyl), 1-(3-hydroxy-pyrrolidinyl), 1-(3-(S)-hydroxy-pyrrolidinyl), 1-piperidinyl, 1-(3-ethoxycarbonyl-piperidinyl), 1-(4-ethoxycarbonyl-piperidinyl), 1-imidazolyl, 1-(2-(phenylamino-methyl)-N-pyrrolidinyl), 1-(3-(R)-dimethylamino-pyrrolidinyl), 1-(3-(R)-hydroxy-pyrrolidinyl), 1-(3,4-dihydroxy-2,5-bis-hydroxymethyl-pyrrolidinyl), 1-(3-(R)-t-butoxycarbonylamino-pyrrolidinyl), 1-(3-(S)-ethylamino-pyrrolidinyl), 1-(3-(R)-amino-pyrrolidinyl), 1-(3-(S)-amino-pyrrolidinyl), 1-(3-(R)-methylamino-pyrrolidinyl), 1-(3-(S)-methylamino-pyrrolidinyl), 1-(3-(N-methyl-N-t-butoxycarbonyl-amino)-pyrrolidinyl) or 1-(2-(3,5-dichlorophenyl)-3-methyl-5-carboxy-1,2,4-triazolyl);

R³ is selected from the group consisting of phenyl and 4-fluorophenyl;

L¹ is selected from the group consisting of -CH₂- , -CH(CH₃)- and -CH₂CH₂-;



is selected from the group consisting of cyclooctyl, 1-acenaphthienyl, R-1-acenaphthienyl, S-1-acenaphthienyl, cyclohexyl, phenyl, 1-naphthyl, and 2-naphthyl,

~~1,2,3,4-tetrahydro-naphthyl, 2-thienyl, benzothienyl, 4,5,6,7-tetrahydro-benzothienyl, bicyclo[3.1.1]hepten-2-yl, bicyclo[3.1.1]heptyl and (3aS)-2,3,3a,4,5,6-hexahydro-1H-phenalen-1-yl;~~

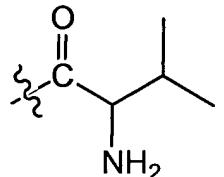
R⁵ is selected from the group consisting of chloro, methyl, n-propyl and trifluoromethyl;

R⁷ is selected from the group consisting of phenyl and 2-thienyl; or a pharmaceutically acceptable salt thereof.

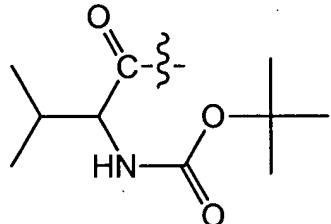
5. (Currently Amended) A compound as in Claim 4 wherein

X is ~~selected from the group consisting of -NR¹R², -SR¹ and -S-CH₂CH(CO₂H)-NH-C(O)-CH₃-~~

R¹ is selected from the group consisting of hydrogen, methyl, ethyl, n-propyl, n-



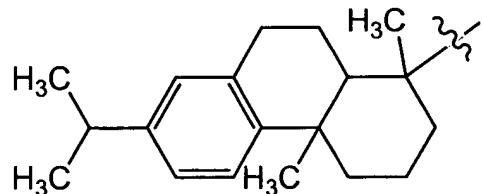
butyl, t-butyl, dimethylaminoethyl, benzyl, phenylethyl,



, 3-methyl-phenyl, 2-(3,4-dimethoxyphenyl)-ethyl, ethoxycarbonyl-methyl, dimethylamino-ethyl and 2-amino-2-methoxycarbonyl-ethyl;

R² is selected from the group consisting of hydrogen, methyl, methoxy, ethyl, ethoxycarbonyl-methyl, 2,2,2-trifluoroethyl, ethoxy, dimethylaminoethyl, n-butyl, t-butyl, n-propyl, di(n-butyl)amino-n-propyl, 3-phenyl-n-propyl, 3-(2-pyridyl)-n-propyl, cyclopropyl, phenyl, 4-fluorophenyl, 4-methylphenyl, 2-aminophenyl, 4-(t-butoxycarbonylamino-ethyl)-phenyl, 3,4-dimethoxyphenyl, 4-biphenyl, 2-ethoxyphenyl, 4-((1-phenyl-pyrazol-2-yl)-amino-sulfonyl)-phenyl, 4-(aminoethyl)-phenyl, benzyl, benzyloxy, 2-methylbenzyl, 3-methylbenzyl, 4-methylbenzyl, 2-methoxybenzyl, 3-methoxybenzyl, 4-methoxybenzyl, 2-ethoxybenzyl, 3-ethoxybenzyl, 2-bromobenzyl, 3-bromobenzyl, 4-bromobenzyl, 3-chlorobenzyl, 4-chlorobenzyl, 3-iodobenzyl, 2-

fluorobenzyl, 3-fluorobenzyl, 4-fluorobenzyl, 2-trifluoromethylbenzyl, 3-trifluoromethylbenzyl, 4-trifluoromethylbenzyl, 4-trifluoromethoxybenzyl, 4-methoxycarbonyl-benzyl, 2,3-dimethoxybenzyl, 2,4-dichlorobenzyl, 3,4-dichlorobenzyl, 2,4-difluorobenzyl, 2,5-difluorobenzyl, 3,4-difluorobenzyl, 3,4,5-trimethoxybenzyl, 2,4,6-trimethoxybenzyl, 4-carboxybenzyl, 3-nitrobenzyl, 4-nitrobenzyl, 2,4-dimethoxybenzyl, 3,4-dimethoxybenzyl, 3,5-dimethoxybenzyl, 3,4-difluorobenzyl, 3,5-di(trifluoromethyl)-benzyl, 2-phenylethyl, 2-(4-bromophenyl)ethyl, 2-(3-methoxyphenyl)ethyl, 2-(4-methoxyphenyl)ethyl, 2-(3,4-dimethoxyphenyl)ethyl, 2-(2-nitro-4,5-dimethoxy-phenyl)ethyl, 3-(4-morpholinyl)-n-propyl, 2-(4-morpholinyl)ethyl, 2-(4-imidazolyl)ethyl, adamantanyl, 1-adamantanyl-methyl, 2-(2,5-dimethoxy-2,5-dihydro-furyl)methyl, 2-pyridyl, 3-pyridyl, 4-pyridyl-methyl, 3-pyridyl-methyl, 4-pyridyl-methyl, 2-(3,4-dimethyl-pyridyl), 2-(5-bromopyridyl), 2-(4,6-dimethyl-pyridyl), 2-(5-methyl-pyridyl), 3-(6-methoxy-pyridyl), thienylmethyl, 2-thienylethyl, 1-naphthyl, 1-naphthyl-methyl, 1-(3,4-methylenedioxyphe-nyl)methyl, 2-(3,4-methylenedioxyphe-nyl)ethyl, 2-furyl-methyl,



, 2S-hydroxy-S-cyclopentyl-methyl, 2S-hydroxy-S-cyclohexyl-methyl, 2S-hydroxy-S-cycloheptyl-methyl; and 2-phenoxy-ethyl and 2-(6-fluoro-2-indetyl)-ethyl;

alternatively when R¹ and R² are both bound to the same nitrogen atom, R¹ and R² are taken together with the nitrogen atom to which they are bound to form a group selected from 1-(4-(3-trifluoromethyl-phenyl)piperazinyl), 1-(4-phenyl-piperidinyl), 1-(4-piperidinyl-piperidinyl), 1-(4-(3,4-methylenedioxyphe-nyl)-methyl)piperazinyl), 1-(3-diethylaminocarbonyl)piperidinyl), 1-(4-[(4-chlorophenyl)phenylmethyl]piperazinyl), 2-(1,2,3,4-tetrahydro-6,7-dimethoxy-isoquinolinyl), 1-(4-t-butoxycarbonyl)piperazinyl), 2-(1,2,3,4-tetrahydro-6,7-dimethoxy-isoquinolinyl), 4-(2,6-dimethyl-morpholinyl), 1-(4-benzyl-piperazinyl), 1-morpholinyl, 1-pyrrolidinyl, 1-(2,3-dihydro-pyrrolidinyl), 1-piperidinyl, 1-(3,5-dimethyl-piperidinyl), 1-(3-hydroxymethyl-piperidinyl), 1-(3-ethoxycarbonyl-

~~piperidinyl), 1-(4-(ethoxycarbonyl)-piperidinyl), 1-imidazolyl and 1-(2-(phenylamino-methyl)-N-pyrrolidinyl);~~

L^1 is selected from the group consisting of $-CH_2-$ and $-CH_2-CH_2-$;



is selected from the group consisting of cyclooctyl, 1-acenaphthenyl, R-1-acenaphthenyl, S-1-acenaphthenyl, cyclohexyl, phenyl, and 1-naphthyl and (3a-S)-2,3,3a,4,5,6-hexahydro-1H-phenalen-2-yl;

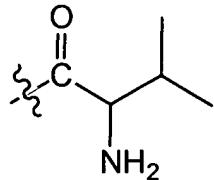
p is an integer from 0 to 2;

R^7 is 2-thienyl;

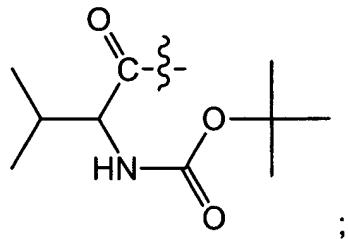
or a pharmaceutically acceptable salt thereof.

6. (Currently Amended) A compound as in Claim 5 wherein

R^1 is selected from the group consisting of hydrogen, methyl, ethyl, n-propyl, n-butyl, t-butyl, dimethylaminoethyl, benzyl, phenylethyl, 2-(3,4-dimethoxyphenyl)-ethyl,

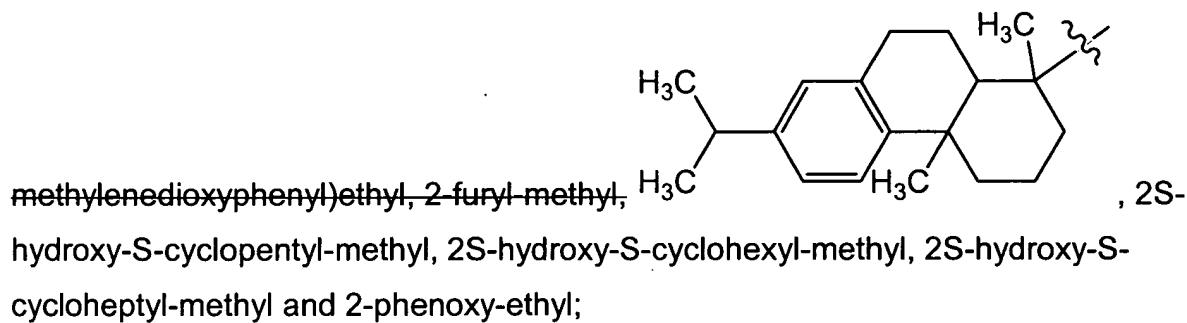


dimethylamino-ethyl, ethoxycarbonyl-methyl, and



R^2 is selected from the group consisting of hydrogen, methyl, methoxy, ethyl, ethoxycarbonyl-methyl, ethoxy, dimethylaminoethyl, n-butyl, n-propyl, di(n-butyl)amino-n-propyl, 3-phenyl-n-propyl, 3-(2-pyridyl)-n-propyl, cyclopropyl, phenyl, 4-fluorophenyl, 4-methylphenyl, 2-aminophenyl, 3,4-dimethoxyphenyl, 4-(t-butoxycarbonylamino-ethyl)-phenyl, 4-biphenyl, 2-ethoxyphenyl, 4-((1-phenyl-pyrazol-2-yl)-aminoсуfonyl)-phenyl, 4-(aminoethyl)-phenyl, benzyl, benzyloxy, 2-methylbenzyl, 3-methylbenzyl, 4-methylbenzyl, 2-methoxybenzyl, 3-methoxybenzyl, 4-methoxybenzyl, 2-ethoxybenzyl, 3-ethoxybenzyl, 2-bromobenzyl, 3-bromobenzyl, 4-bromobenzyl, 3-chlorobenzyl, 4-

chlorobenzyl, 3-iodobenzyl, 2-fluorobenzyl, 3-fluorobenzyl, 4-fluorobenzyl, 2-trifluoromethylbenzyl, 3-trifluoromethylbenzyl, 4-trifluoromethylbenzyl, 4-trifluoromethoxybenzyl, 4-methoxycarbonylbenzyl, 2,3-dimethoxybenzyl, 2,4-dichlorobenzyl, 3,4-dichlorobenzyl, 2,4-difluorobenzyl, 2,5-difluorobenzyl, 3,4,5-trimethoxybenzyl, 2,4,6-trimethoxybenzyl, 3-nitrobenzyl, 4-nitrobenzyl, 2,4-dimethoxybenzyl, 3,4-dimethoxybenzyl, 3,5-dimethoxybenzyl, 3,4-difluorobenzyl, 3,5-di(trifluoromethyl)-benzyl, 2-phenylethyl, 2-(4-bromophenyl)ethyl, 2-(3-methoxyphenyl)ethyl, 2-(4-methoxyphenyl)ethyl, 2-(3,4-dimethoxyphenyl)ethyl, 2-(2-nitro-4,5-dimethoxy-phenyl)ethyl, 3-(4-morpholinyl)-n-propyl, 2-(4-morpholinyl)ethyl, 2-(4-imidazolyl)ethyl, 1-adamantanyl, 1-adamantanyl-methyl, 2-(2,5-dimethoxy-2,5-dihydro-furyl)methyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, 2-pyridyl-methyl, 3-pyridyl-methyl, 4-pyridyl-methyl, 2-(3,4-dimethyl-pyridyl), 2-(5-bromopyridyl), 2-(4,6-dimethyl-pyridyl), 2-(5-methyl-pyridyl), 3-(6-methoxy-pyridyl), thienylmethyl, 2-thienylethyl, 1-naphthyl, 1-naphthyl-methyl, 1-(3,4-methylenedioxophenyl)methyl, 2-(3,4-



alternatively when R¹ and R² are both bound to the same nitrogen atom, R¹ and R² are taken together with the nitrogen atom to which they are bound to form a group selected from 1-(4-(3-trifluoromethyl-phenyl)piperazinyl), 1-(4-phenyl-piperidinyl), 1-(4-piperidinyl-piperidinyl), 1-(4-(3,4-methylenedioxophenyl-methyl)piperazinyl), 1-(3-diethylaminocarbonyl)piperidinyl), 1-(4-[(4-chlorophenyl)phenylmethyl]piperazinyl), 2-(1,2,3,4-tetrahydro-6,7-dimethoxy-isoquinolinyl), 1-(4-t-butoxycarbonyl)piperazinyl), 2-(1,2,3,4-tetrahydro-6,7-dimethoxy-isoquinolinyl), 4-(2,6-dimethyl-morpholinyl), 1-(4-benzyl-piperazinyl), 1-(3,5-dimethyl-piperidinyl), 1-(3-hydroxymethyl-piperidinyl), 1-(3-ethoxycarbonyl-piperidinyl), 1-(4-ethoxycarbonyl)piperidinyl), 1-piperidinyl, 1-

~~morpholinyl, 1-pyrrolidinyl, 1-imidazolyl, 1-(2,3-dihydro-pyrrolidinyl), and 1-(2-(phenylamino-methyl)-N-pyrrolidinyl);~~

p is an integer from 0 to 1;

R⁵ is selected from the group consisting of methyl, n-propyl and trifluoromethyl; or a pharmaceutically acceptable salt thereof.

7. (Currently Amended) A compound as in Claim 4 wherein

R⁰ is -CH₂-CH(OH)-CH₂-X;

X is-NR¹R²;

R¹ is selected from the group consisting of hydrogen, 2-(3,4-dimethoxyphenyl)-ethyl, 1-(3,4-dimethoxyphenyl)-n-ethyl and amino-n-propyl;

R² is selected from the group consisting of hydrogen, methyl, n-butyl, 3-hydroxy-n-propyl, 3-methoxy-n-propyl, methylamino-n-propyl, dimethylamino-n-propyl, t-butoxycarbonylamino-n-propyl, N-methyl-N-t-butoxycarbonyl-amino-n-ethyl, 3-nitrobenzyl, 4-methoxycarbonyl-benzyl, and -CH(CH₃)-phenyl, 4-pyridinyl, 1-(4-ethoxycarbonyl-piperidinyl) and 2-(3H-imidazol-4-yl)-ethyl;

~~alternatively when R¹ and R² are both bound to the same nitrogen atom, R¹ and R² are taken together with the nitrogen atom to which they are bound to form a group selected from 2-(1,2,3,4-tetrahydro-6,7-dimethoxy-isquinolinyl), 1-(4-[(4-chlorophenyl)-phenyl-methyl]piperazinyl), 1-pyrrolidinyl, 1-(3-hydroxy-pyrrolidinyl), 1-(3-(S)-hydroxy-pyrrolidinyl), 1-(3-(R)-hydroxy-pyrrolidinyl), 1-(4-hydroxy-piperidinyl), 1-(3-(R)-dimethylamino-pyrrolidinyl), 1-(4-t-butoxycarbonylamino-pyrrolidinyl), 1-(3-(R)-t-butoxycarbonylamino-pyrrolidinyl), 1-(3-(R)-amino-pyrrolidinyl), 1-(3-(S)-amino-pyrrolidinyl), 1-(3-(R)-methylamino-pyrrolidinyl), 1-(3-(S)-methylamino-pyrrolidinyl), 1-(3-(S)-ethylamino-pyrrolidinyl), 1-(4-dimethylamino-pyrrolidinyl), 1-(3-(N-methyl-N-t-butoxycarbonyl-amino-pyrrolidinyl) or 1-(2-(3,5-dichlorophenyl)-3-methyl-5-carboxy-1,2,4-triazolyl);~~

R³ is selected from the group consisting of phenyl and 4-fluorophenyl;

L¹ is selected from the group consisting of -CH₂- and -CH₂CH₂-;



is selected from the group consisting cyclooctyl, 1-naphthyl, 1-acenaphthenyl, R-1-acenaphthenyl, and S-1-acenaphthenyl, bicyclo[3.1.1.]hepten-2-yl, bicyclo[3.1.1.]heptyl and (3aS)-2,3,3a,4,5,6-hexahydro-1H-phenalen-1-yl;.

p is an integer from 0 to 1;

R⁵ is methyl;

q is 0;

or a pharmaceutically acceptable salt thereof.

8. (Currently Amended) A compound as in Claim 7 wherein

R¹ is selected from the group consisting of hydrogen, 1-(3,4-dimethoxyphenyl)-n-ethyl and amino-n-propyl;

R² is selected from the group consisting of hydrogen, methyl, n-butyl, 3-hydroxy-n-propyl, 3-methoxy-n-propyl, methylamino-n-propyl, dimethylamino-n-propyl, N-methyl-N-t-butoxycarbonyl-amino-n-ethyl, 3-nitrobenzyl, 4-methoxycarbonyl-benzyl, and -CH(CH₃)-phenyl, 4-pyridinyl and 2-(3H-imidazol-4-yl)-ethyl;

alternatively when R¹ and R² are both bound to the same nitrogen atom, R¹ and R² are taken together with the nitrogen atom to which they are bound to form a group selected from 2-(1,2,3,4-tetrahydro-6,7-dimethoxy-isquinolinyl), 1-(4-[(4-chlorophenyl)-phenyl-methyl]piperazinyl), 1-pyrrolidinyl, 1-(3-hydroxy-pyrrolidinyl), 1-(3-(R)-hydroxy-pyrrolidinyl), 1-(4-hydroxy-piperidinyl), 1-(3-(R)-dimethylamino-pyrrolidinyl), 1-(4-t-butoxycarbonylamino-pyrrolidinyl), 1-(3-(R)-t-butoxycarbonylamino-pyrrolidinyl), 1-(3-(R)-amino-pyrrolidinyl), 1-(3-(S)-amino-pyrrolidinyl), 1-(3-(R)-methylamino-pyrrolidinyl), 1-(3-(S)-methyamino-pyrrolidinyl), 1-(3-(S)-ethylamino-pyrrolidinyl), 1-(4-dimethylamino-pyrrolidinyl), 1-(3-(N-methyl-N-t-butoxycarbonyl-amino-pyrrolidinyl)) or 1-(2-(3,5-dichlorophenyl)-3-methyl-5-carboxy-1,2,4-triazolyl);



is selected from the group consisting cyclooctyl, 1-naphthyl, 1-acenaphthenyl, R-1-acenaphthenyl, and S-1-acenaphthenyl, bicyclo[3.1.1.]hepten-2-yl, bicyclo[3.1.1.]heptyl and (3aS)-2,3,3a,4,5,6-hexahydro-1H-phenalen-1-yl;.

or a pharmaceutically acceptable salt thereof.

9. (Currently Amended) A compound as in Claim 1 selected from the group consisting of

8-(R) acenaphthen-1-yl-3-(3-amino-2-(S)-hydroxy-propyl)-1-(4-fluoro-phenyl)-1,3,8-triaza-spiro[4.5]decan-4-one;

8-(R) acenaphthen-1-yl-3-(3-amino-2-(R)-hydroxy-propyl)-1-(4-fluoro-phenyl)-1,3,8-triaza-spiro[4.5]decan-4-one;

8-(R)-Acenaphthen-1-yl-3-(3-dimethylamino-2-(R)-hydroxy-propyl)-1-(4-fluoro-phenyl)-1,3,8-triaza-spiro[4.5]decan-4-one;

3-(3-Amino-2-(R)-hydroxy-propyl)-1-(4-fluoro-phenyl)-8-(8-methyl-naphthalen-1-ylmethyl)-1,3,8-triaza-spiro[4.5]decan-4-one;

3-(3-Dimethylamino-2-(R)-hydroxy-propyl)-1-(4-fluoro-phenyl)-8-(8-methyl-naphthalen-1-ylmethyl)-1,3,8-triaza-spiro[4.5]decan-4-one;

~~8-(R) Acenaphthen-1-yl-1-(4-fluoro-phenyl)-3-[2-(R)-hydroxy-3-(3-hydroxymethyl-piperidin-1-yl)-propyl]-1,3,8-triaza-spiro[4.5]decan-4-one;~~

~~3-(3-Amino-2-(R)-hydroxy-propyl)-8-cyclooctyl-1-(4-fluoro-phenyl)-1,3,8-triaza-spiro[4.5]decan-4-one;~~

~~3-(3-Amino-2-(R)-hydroxy-propyl)-1-(4-fluoro-phenyl)-8-1-(S)-(3aS)-(2,3,3a,4,5,6-hexahydro-1H-phenalen-1-yl)-1,3,8-triaza-spiro[4.5]decan-4-one;~~

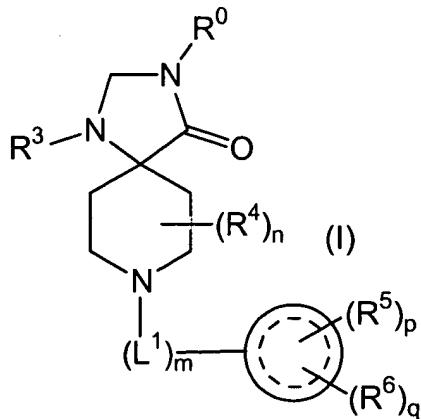
1-(4-Fluoro-phenyl)-3-[2-(R)-hydroxy-3-(3-hydroxy-propylamino)-propyl]-8-(8-methyl-naphthalen-1-ylmethyl)-1,3,8-triaza-spiro[4.5]decan-4-one;

1-(4-Fluoro-phenyl)-3-[2-(R)-hydroxy-3-(3-methylamino-propylamino)-propyl]-8-(8-methyl-naphthalen-1-ylmethyl)-1,3,8-triaza-spiro[4.5]decan-4-one;

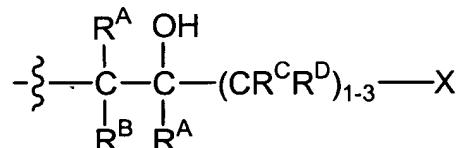
3-[3-(3-Dimethylamino-propylamino)-2-(R)-hydroxy-propyl]-1-(4-fluoro-phenyl)-8-(8-methyl-naphthalen-1-ylmethyl)-1,3,8-triaza-spiro[4.5]decan-4-one

and pharmaceutically acceptable salts thereof.

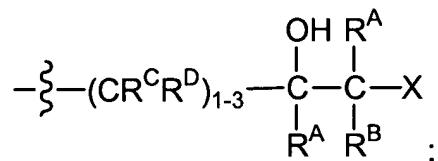
10. (Currently Amended) A compound of the formula (I)



wherein



R^0 is selected from the group consisting of and



each R^{A} and R^{B} is independently selected from the group consisting of hydrogen and $\text{C}_{1-4}\text{alkyl}$;

each R^{C} and R^{D} is independently selected from the group consisting of hydrogen, hydroxy, carboxy, $\text{C}_{1-4}\text{alkyl}$, $\text{C}_{1-4}\text{alkoxy}$, nitro, cyano, $\text{N}(\text{R}^{\text{E}})_2$, aryl, ar $\text{C}_{1-4}\text{alkyl}$, heteroaryl or heterocycloalkyl; wherein the aryl, ar $\text{C}_{1-4}\text{alkyl}$, heteroaryl or heterocycloalkyl substituent is optionally substituted with one or more substituents independently selected from hydroxy, carboxy, $\text{C}_{1-4}\text{alkyl}$, $\text{C}_{1-4}\text{alkoxy}$, nitro, cyano or $\text{N}(\text{R}^{\text{E}})_2$;

each R^{E} is independently selected from the group consisting of hydrogen and $\text{C}_{1-4}\text{alkyl}$;

X is selected from the group consisting of $-\text{NR}^1\text{R}^2$, $-\text{C}(\text{O})-\text{NR}^1\text{R}^2$, $-\text{NR}^1-\text{C}(\text{O})-\text{R}^2$, $-\text{OR}^1$, $-\text{SR}^1$, $-\text{SOR}^1$, $-\text{SO}_2\text{R}^1$, $-\text{S}-(\text{C}_{2-4}\text{alkyl})-\text{NR}^1\text{R}^2$, $-\text{S}-(\text{C}_{2-4}\text{alkyl})-\text{C}(\text{O})\text{O}-\text{C}(\text{CH}_3)_3$, $-\text{SO}-(\text{C}_{1-4}\text{alkyl})-\text{NR}^1\text{R}^2$ and $-\text{SO}_2-(\text{C}_{1-4}\text{alkyl})-\text{NR}^1\text{R}^2$; wherein the alkyl portion of the $-\text{S}-(\text{C}_{2-4}\text{alkyl})-\text{NR}^1\text{R}^2$, $-\text{SO}-(\text{C}_{1-4}\text{alkyl})-\text{NR}^1\text{R}^2$ or $-\text{SO}_2-(\text{C}_{1-4}\text{alkyl})-\text{NR}^1\text{R}^2$ group is optionally substituted with one or more substituents independently selected from carboxy, hydroxy, hydroxy $\text{C}_{1-4}\text{alkyl}$, $\text{C}_{1-4}\text{alkyl}$, $\text{C}_{1-4}\text{alkoxycarbonyl}$ or $-\text{CONR}^1\text{R}^2$;

each R¹ and R² is independently selected from the group consisting of hydrogen, C₁₋₈alkyl, C₁₋₈alkoxy, cycloalkyl, cycloalkyl-C₁₋₄alkyl, partially unsaturated carbocyclyl, aryl, arC₁₋₄alkyl, arC₁₋₄alkoxy, heteroaryl, heteroaryl-C₁₋₄alkyl, heterocycloalkyl, heterocycloalkyl-C₁₋₄alkyl, -C(O)-C₁₋₆alkyl, -C(O)-aryl, and -C(O)-arC₁₋₄alkyl, -C(O)-heteroaryl and -C(O)-heterocycloalkyl; wherein the C₁₋₈alkyl, cycloalkyl, partially unsaturated carbocyclyl, aryl, or arC₁₋₈alkyl, heteroaryl or heterocycloalkyl group, whether alone or part of a substituent group, is optionally substituted with one or more substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, trifluoromethyl, trifluoromethoxy, nitro, cyano, -C(O)-C₁₋₄alkyl, C₁₋₄alkoxycarbonyl, N(R^E)₂, N(R^E)₂-C₁₋₄alkyl, N(R^E)-C(O)C(CH₃)₃, aryl, aryloxy, cycloalkyl, heteroaryl, aryl substituted heteroarylaminosulfonyl or C₁₋₆alkylthio;

alternatively when R¹ and R² are both bound to the same nitrogen atom, R¹ and R² are taken together with the nitrogen atom to which they are bound to form a heteroaryl or heterocycloalkyl group; wherein the heteroaryl or heterocycloalkyl group is optionally substituted with one or more substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, C₁₋₄alkoxycarbonyl, trifluoromethyl, trifluoromethoxy, nitro, cyano, N(R^E)₂, aryl, arC₁₋₄alkyl, heteroaryl, heterocycloalkyl, di(C₁₋₆)alkylamine-carbonyl, t-butoxycarbonyl or arylamino-C₁₋₄alkyl; wherein the aryl, arC₁₋₄alkyl, heteroaryl or heterocycloalkyl substituent is optionally further substituted with one or more substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, trifluoromethyl, trifluoromethoxy, nitro, cyano, N(R^E)₂ or substituted phenyl; wherein the substituents on the phenyl are one or more independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, trifluoromethyl, trifluoromethoxy, nitro, cyano or N(R^E)₂;

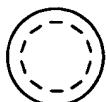
R³ is selected from the group consisting of aryl, arC₁₋₆alkyl and heteroaryl; wherein the aryl, arC₁₋₆alkyl or heteroaryl group is optionally substituted with one or more substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, trifluoromethyl, trifluoromethoxy, nitro, cyano or N(R^E)₂;

n is an integer from 0 to 2;

R⁴ is selected from the group consisting of hydroxy, C₁₋₄alkyl and hydroxy substituted C₁₋₄alkyl;

m is an integer from 0 to 1;

L¹ is selected from the group consisting of C₁₋₆alkyl and C₃₋₆alkenyl; wherein the double bond of the C₃₋₆alkenyl group is at least one carbon atom removed from the attachment point to the N atom; and wherein the C₁₋₆alkyl or C₃₋₆alkenyl group is optionally substituted with one to two substituents independently selected from hydroxy, fluoro, C₁₋₆alkyl, fluorinated C₁₋₆alkyl or C₁₋₆alkoxy;



is selected from the group consisting of ~~cycloalkyl, partially unsaturated carbocyclyl, aryl, heteroaryl and heterocycloalkyl~~ phenyl, naphthyl and acenaphthyl;

p is an integer from 0 to 5;

R⁵ is selected from the group consisting of hydroxy, carboxy, halogen, C₁₋₆alkyl, C₁₋₆alkoxy, nitro, cyano, NR¹R², trifluoromethyl, trifluoromethoxy, C₁₋₄alkoxycarbonyl, -SO-NR¹R², -SO₂-NR¹R² and -C(O)-NR¹R²,

q is an integer from 0 to 1;

R⁶ is selected from the group consisting of -(L²)₀₋₁-R⁷;

L² is selected from the group consisting of -C₁₋₆alkyl-, -C₂₋₄alkenyl-, -C₂₋₆alkynyl-, -O-, -S-, -NH-, -N(C₁₋₄alkyl)-, -C₁₋₆alkyl-O-, -C₁₋₆alkyl-S-, -O-C₁₋₆alkyl-, -S-C₁₋₆alkyl-, -O-C₂₋₆alkyl-O-, -S-C₂₋₆alkyl-S-, -SO₂-, -SO₂NH-, -SO₂N(C₁₋₄alkyl)-, -NH-SO₂-, -N(C₁₋₄alkyl)-SO₂-, -C(O)-O- and -O-C(O)-;

R⁷ is selected from the group consisting of aryl, partially unsaturated carbocyclyl, cycloalkyl, heteroaryl and heterocycloalkyl; wherein the aryl, partially unsaturated carbocyclyl, cycloalkyl, heteroaryl or heterocycloalkyl group is optionally substituted with one or more substituents independently selected from hydroxy, carboxy, halogen, C₁₋₆alkyl, C₁₋₆alkoxy, nitro, cyano, N(R^E)₂, trifluoromethyl, trifluoromethoxy, C₁₋₄alkoxycarbonyl, -SO₂-N(R^E)₂ and -C(O)-N(R^E)₂;

or a pharmaceutically acceptable salt thereof.

11. (Original) A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a compound of Claim 1.

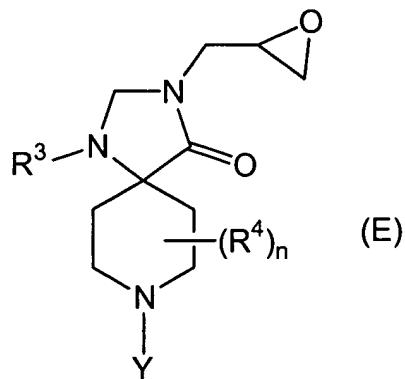
12. (Original) A pharmaceutical composition made by mixing a compound of Claim 1 and a pharmaceutically acceptable carrier.
13. (Original) A process for making a pharmaceutical composition comprising mixing a compound of Claim 1 and a pharmaceutically acceptable carrier.
14. (Withdrawn) A method of treating a disorder mediated by the ORL-1 receptor, in a subject in need thereof comprising administering to the subject a therapeutically effective amount of the compound of Claim 1.
15. (Withdrawn) The method of Claim 14, wherein the disorder mediated by the ORL-1 receptor is selected from the group consisting of anxiety, depression, panic, mania, dementia, bipolar disorder, substance abuse, neuropathic pain, acute pain, chronic pain, migraine, asthma, cough, psychosis, schizophrenia, epilepsy, hypertension, obesity, eating disorders, cravings, diabetes, cardiac arrhythmia, irritable bowel syndrome, Crohn's disease, urinary incontinence, adrenal disorders, attention deficit disorder (ADD), attention deficit hyperactivity disorder (ADHD), Alzheimer's disease, improved cognition, improved memory and mood stabilization.
16. (Withdrawn) A method of treating a disorder mediated by the ORL-1 receptor, in a subject in need thereof comprising administering to the subject a therapeutically effective amount of the composition of Claim 11.
17. (Withdrawn) A method of treating a condition selected from the group consisting of anxiety, depression, panic, mania, dementia, bipolar disorder, substance abuse, neuropathic pain, acute pain, chronic pain, migraine, asthma, cough, psychosis, schizophrenia, epilepsy, hypertension, obesity, eating disorders, cravings, diabetes, cardiac arrhythmia, irritable bowel syndrome, Crohn's disease, urinary incontinence, adrenal disorders, attention deficit disorder (ADD), attention deficit hyperactivity disorder (ADHD), Alzheimer's disease, improved cognition, improved memory and mood

stabilization, in a subject in need thereof comprising administering to the subject a therapeutically effective amount of the compound of Claim 1.

18. (Withdrawn) A method of treating a condition selected from the group consisting of anxiety, depression, panic, mania, dementia, bipolar disorder, substance abuse, neuropathic pain, acute pain, chronic pain, migraine, asthma, cough, psychosis, schizophrenia, epilepsy, hypertension, obesity, eating disorders, cravings, diabetes, cardiac arrhythmia, irritable bowel syndrome, Crohn's disease, urinary incontinence, adrenal disorders, attention deficit disorder (ADD), attention deficit hyperactivity disorder (ADHD), Alzheimer's disease, improved cognition, improved memory and mood stabilization, in a subject in need thereof comprising administering to the subject a therapeutically effective amount of the composition of Claim 7.

19. (Withdrawn) The use of a compound as in Claim 1 for the preparation of a medicament for the treatment of (a) anxiety, (b) depression, (c) panic, (d) mania, (e) dementia, (f) bipolar disorder, (g) substance abuse, (h) neuropathic pain, (i) acute pain, (j) chronic pain, (k) migraine, (l) asthma, (m) cough, (n) psychosis, (o) schizophrenia, (p) epilepsy, (q) hypertension, (r) obesity, (s) eating disorders, (t) cravings, (u) diabetes, (v) cardiac arrhythmia, (w) irritable bowel syndrome, (x) Crohn's disease, (y) urinary incontinence, (z) adrenal disorders, (aa) attention deficit disorder (ADD), (bb) attention deficit hyperactivity disorder (ADHD), (cc) Alzheimer's disease, for (dd) improved cognition, (ee) improved memory or (ff) mood stabilization, in a subject in need thereof.

20. (Withdrawn) A compound of the formula (E)



wherein

R^3 is selected from the group consisting of aryl, arC₁₋₆alkyl and heteroaryl; wherein the aryl, arC₁₋₆alkyl or heteroaryl group is optionally substituted with one or more substituents independently selected from halogen, hydroxy, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, trifluoromethyl, trifluoromethoxy, nitro, cyano or N(R^E)₂;

each R^E is independently selected from hydrogen or C₁₋₄alkyl;

n is an integer from 0 to 2;

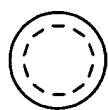
R^4 is selected from the group consisting of hydroxy, C₁₋₄alkyl and hydroxy substituted C₁₋₄alkyl;

Y is selected from the group consisting of hydrogen, C₁₋₄alkyl, t-butoxycarbonyl

$\begin{array}{c} \text{---} \\ | \\ -\xi-(L^1)_m-\text{---} \\ | \\ \text{---} \end{array}$ (R⁵)_p
and (R⁶)_q;

m is an integer from 0 to 1;

L^1 is selected from the group consisting of C₁₋₆alkyl and C₃₋₆alkenyl; wherein the double bond of the C₃₋₆alkenyl group is at least one carbon atom removed from the attachment point to the N atom; and wherein the C₁₋₆alkyl or C₃₋₆alkenyl group is optionally substituted with one to two substituents independently selected from hydroxy, fluoro, C₁₋₆alkyl, fluorinated C₁₋₆alkyl or C₁₋₆alkoxy;



is selected from the group consisting of cycloalkyl, partially unsaturated carbocyclyl, aryl, heteroaryl and heterocycloalkyl;

p is an integer from 0 to 5;

R^5 is selected from the group consisting of hydroxy, carboxy, halogen, C_{1-6} alkyl, hydroxy substituted C_{1-6} alkyl, C_{1-6} alkoxy, nitro, cyano, NR^1R^2 , trifluoromethyl, trifluoromethoxy, C_{1-4} alkoxycarbonyl, $-SO-NR^1R^2$, $-SO_2-NR^1R^2$ and $-C(O)-NR^1R^2$;

q is an integer from 0 to 1;

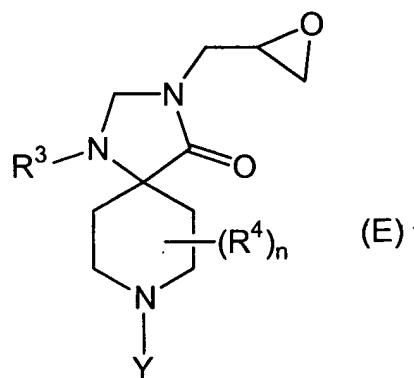
R^6 is selected from the group consisting of $-(L^2)_{0-1}R^7$;

L^2 is selected from the group consisting of $-C_{1-6}$ alkyl-, $-C_{2-4}$ alkenyl-, $-C_{2-6}$ alkynyl-, $-O-$, $-S-$, $-NH-$, $-N(C_{1-4}$ alkyl)-, $-C_{1-6}$ alkyl-O-, $-C_{1-6}$ alkyl-S-, $-O-C_{1-6}$ alkyl-, $-S-C_{1-6}$ alkyl-, $-O-C_{2-6}$ alkyl-O-, $-S-C_{2-6}$ alkyl-S-, $-SO_2-$, $-SO_2NH-$, $-SO_2N(C_{1-4}$ alkyl)-, $-NH-SO_2-$, $-N(C_{1-4}$ alkyl)- SO_2- , $-C(O)-O-$ and $-O-C(O)-$;

R^7 is selected from the group consisting of aryl, partially unsaturated carbocyclyl, cycloalkyl, heteroaryl and heterocycloalkyl; wherein the aryl, partially unsaturated carbocyclyl, cycloalkyl, heteroaryl or heterocycloalkyl group is optionally substituted with one or more substituents independently selected from hydroxy, carboxy, halogen, C_{1-6} alkyl, C_{1-6} alkoxy, nitro, cyano, $N(R^E)_2$, trifluoromethyl, trifluoromethoxy, C_{1-4} alkoxycarbonyl, $-SO_2-N(R^E)_2$ and $-C(O)-N(R^E)_2$;

or a pharmaceutically acceptable salt thereof.

21. (Withdrawn) A compound of the formula (E)



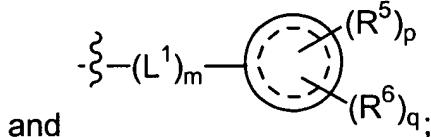
wherein

R^3 is selected from the group consisting of aryl, arC_{1-6} alkyl and heteroaryl; wherein the aryl, arC_{1-6} alkyl or heteroaryl group is optionally substituted with one or more substituents independently selected from halogen, hydroxy, carboxy, C_{1-4} alkyl, C_{1-4} alkoxy, trifluoromethyl, trifluoromethoxy, nitro, cyano or $N(R^E)_2$;

each R^E is independently selected from hydrogen or C₁₋₄alkyl;
n is an integer from 0 to 2;

R⁴ is selected from the group consisting of hydroxy, C₁₋₄alkyl and hydroxy substituted C₁₋₄alkyl;

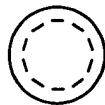
Y is selected from the group consisting of hydrogen, C₁₋₄alkyl, t-butoxycarbonyl



and

m is an integer from 0 to 1;

L¹ is selected from the group consisting of C₁₋₆alkyl and C₃₋₆alkenyl; wherein the double bond of the C₃₋₆alkenyl group is at least one carbon atom removed from the attachment point to the N atom; and wherein the C₁₋₆alkyl or C₃₋₆alkenyl group is optionally substituted with one to two substituents independently selected from hydroxy, fluoro, C₁₋₆alkyl, fluorinated C₁₋₆alkyl or C₁₋₆alkoxy;



is selected from the group consisting of cycloalkyl, partially unsaturated carbocyclyl, aryl, heteroaryl and heterocycloalkyl;

p is an integer from 0 to 5;

R⁵ is selected from the group consisting of hydroxy, carboxy, halogen, C₁₋₆alkyl, C₁₋₆alkoxy, nitro, cyano, NR¹R², trifluoromethyl, trifluoromethoxy, C₁₋₄alkoxycarbonyl, -SO-NR¹R², -SO₂-NR¹R² and -C(O)-NR¹R²;

q is an integer from 0 to 1;

R⁶ is selected from the group consisting of -(L²)₀₋₁-R⁷;

L² is selected from the group consisting of -C₁₋₆alkyl-, -C₂₋₄alkenyl-, -C₂₋₆alkynyl-, -O-, -S-, -NH-, -N(C₁₋₄alkyl)-, -C₁₋₆alkyl-O-, -C₁₋₆alkyl-S-, -O-C₁₋₆alkyl-, -S-C₁₋₆alkyl-, -O-C₂₋₆alkyl-O-, -S-C₂₋₆alkyl-S-, -SO₂-, -SO₂NH-, -SO₂N(C₁₋₄alkyl)-, -NH-SO₂-, -N(C₁₋₄alkyl)-SO₂-, -C(O)-O- and -O-C(O)-;

R⁷ is selected from the group consisting of aryl, partially unsaturated carbocyclyl, cycloalkyl, heteroaryl and heterocycloalkyl; wherein the aryl, partially unsaturated carbocyclyl, cycloalkyl, heteroaryl or heterocycloalkyl group is optionally substituted with

one or more substituents independently selected from hydroxy, carboxy, halogen, C₁-₆alkyl, C₁-₆alkoxy, nitro, cyano, N(R^E)₂, trifluoromethyl, trifluoromethoxy, C₁-₄alkoxycarbonyl, -SO₂-N(R^E)₂ and -C(O)-N(R^E)₂;
or a pharmaceutically acceptable salt thereof.